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L1 and (Phe570 or Phe667)	2

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DATE: Tuesday, May 21, 2002 [Printable Copy](#) [Create Case](#)

<u>Set Name</u> side by side	<u>Query</u>	<u>Hit Count</u>	<u>Set Name</u> result set
DB=USPT,PGPB,JPAB,EPAB,DWPI; PLUR=YES; OP=ADJ			
<u>L2</u>	L1 and (Phe570 or Phe667)	2	<u>L2</u>
<u>L1</u>	DNA adj polymerase	18570	<u>L1</u>

END OF SEARCH HISTORY.

WEST[Generate Collection](#)[Print](#)**Search Results - Record(s) 1 through 2 of 2 returned.**☐ 1. Document ID: US 20010012613 A1

L2: Entry 1 of 2

File: PGPB

Aug 9, 2001

PGPUB-DOCUMENT-NUMBER: 20010012613
PGPUB-FILING-TYPE: new
DOCUMENT-IDENTIFIER: US 20010012613 A1

TITLE: THERMOSTABLE POLYMERASES HAVING ALTERED FIDELITY AND METHOD OF IDENTIFYING AND USING SAME

PUBLICATION-DATE: August 9, 2001

INVENTOR-INFORMATION:

NAME	CITY	STATE	COUNTRY	RULE-47
LOEB, LAWRENCE A.	BELLEVUE	WA	US	
HOOD, LEROY	SEATTLE	WA	US	
SUZUKI, MOTOSHI	NOGOYA		JP	

US-CL-CURRENT: 435/6

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWC	Draw Desc	Image
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☐ 2. Document ID: US 5614365 A

L2: Entry 2 of 2

File: USPT

Mar 25, 1997

US-PAT-NO: 5614365
DOCUMENT-IDENTIFIER: US 5614365 A

TITLE: DNA polymerase having modified nucleotide binding site for DNA sequencing

Full	Title	Citation	Front	Review	Classification	Date	Reference	Sequences	Attachments	Claims	KWC	Draw Desc	Image
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Terms	Documents
L1 and (Phe570 or Phe667)	2

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(FILE 'HOME' ENTERED AT 09:58:42 ON 21 MAY 2002)

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BIOBUSINESS, BIOCOMMERCE, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO,
CABA,
CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB,
DDFU, DGENE, DRUGB, DRUGLAUNCH, DRUGMONOG2, ...' ENTERED AT 09:58:55 ON
21 MAY 2002

SEA DNA(W) POLYMERASE

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L1 QUE DNA(W) POLYMERASE

FILE 'MEDLINE, EMBASE, BIOSIS, SCISEARCH, BIOTECHNO, CAPLUS' ENTERED AT
10:04:57 ON 21 MAY 2002

L2 13 S L1 AND (PHE570 OR PHE667) OR (F570 OR F667)
L3 6 DUP REM L2 (7 DUPLICATES REMOVED)
L4 13709 S L1 AND (MUTANT OR VARIANT)
L5 0 S L4 AND (PHE570 OR PHE 570)
L6 38154 S L4 AND (PHE667 OR PHE570) OR (570 OR 667)
L7 3 S L4 AND (PHE667 OR PHE570)
L8 1 DUP REM L7 (2 DUPLICATES REMOVED)

=> d l8 ibib ab

L8 ANSWER 1 OF 1 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.DUPLICATE 1

ACCESSION NUMBER: 2000378102 EMBASE

TITLE: Thermus aquaticus **DNA polymerase I**
mutants with altered fidelity. Interacting
mutations in the O-helix.

AUTHOR: Suzuki M.; Yoshida S.; Adman E.T.; Blank A.; Loeb L.A.;
Gottstein J.

CORPORATE SOURCE: L.A. Loeb, J. Gottstein Mem. Cancer Res. Lab., Dept. of
Pathology, University of Washington, Seattle, WA
98195-7705, United States. laloeb@u.washington.edu

SOURCE: Journal of Biological Chemistry, (20 Oct 2000) 275/42
(32728-32735).

Refs: 35

ISSN: 0021-9258 CODEN: JBCHA3

COUNTRY: United States

DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 029 Clinical Biochemistry

LANGUAGE: English

SUMMARY LANGUAGE: English

AB **Phe667** in the conserved O-helix of *Thermus aquaticus* (Taq)
DNA polymerase I (pol I) is known to be important for
discrimination against dideoxy-NTPs. We show here that **Phe667** is
also important for base selection fidelity. In a forward mutation assay

at
high polymerase concentration, wild type pol I catalyzed frequent A
.fwdarw. T and G .fwdarw. T transversions and -1 frameshifts at
nonreiterated sites involving loss of a purine immediately downstream of

a
pyrimidine. The **mutants** F667L and A661E, I665T, F667L exhibited
large decreases in A .fwdarw. T and G .fwdarw. T transversions, and the
triple **mutant** displayed reduction in the aforementioned -1
frameshifts as well. Kinetic analysis showed that the F667L and
A661E, I665T, F667L polymerases discriminated against synthesis of A:A
mispairs more effectively and catalyzed less extension of A:A mispairs
than the wild type enzyme. These data indicate that **Phe667**
functions in maintaining the error frequency and spectrum, and the
catalytic efficiency, of wild type pol I. We also found that the strong
general mutator activity conferred by the single A661E substitution was
entirely suppressed in the A661E, I665T, F667L polymerase, exemplifying

how
interactions among O-helix residues can contribute to fidelity. We
discuss

the mutator and anti-mutator mutations in light of recently obtained
three-dimensional structures of *T. aquaticus* pol I.

=> d 13 ibib ab 1-6

L3 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2002 ACS
ACCESSION NUMBER: 2000:260305 CAPLUS
DOCUMENT NUMBER: 132:265445
TITLE: Preparation of nucleotide compounds including a rigid linker used in DNA sequencing
INVENTOR(S): Kahn, Shaheer H.; Rosenblum, Barnett B.; Zhen, Weiguo;
Menchen, Steven M.
PATENT ASSIGNEE(S): The Perkin-Elmer Corporation, USA
SOURCE: PCT Int. Appl., 63 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2000021974	A1	20000420	WO 1999-US12323	19990602
W: AU, CA, JP RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
US 6096875	A	20000801	US 1998-172789	19981014
AU 9946740	A1	20000501	AU 1999-46740	19990602
EP 1121371	A1	20010808	EP 1999-930140	19990602
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
PRIORITY APPLN. INFO.:			US 1998-172789	A 19981014
			WO 1999-US12323	W 19990602

OTHER SOURCE(S): MARPAT 132:265445
AB A nucleoside/tide compd. having a rigid linker attached to the 8-position of a purine, the 7-position of a 7-deazapurine and the 5-position of a pyrimidine is disclosed. Fluorescent dyes may be attached to this linker and the fluorescent nucleotide used in primer extension reactions. Thus, the fluorescein dye HEX-1 was attached to the 5-position of ddCTP via an acetylene-phenyl-oxyethyleneamino linkage. This nucleotide deriv. was used in DNA sequencing with Taq polymerase contg. an R660S mutation. The synthesis of a no. of nucleoside/nucleotide derivs. contg. various rigid linkers is described.
REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L3 ANSWER 2 OF 6 EMBASE COPYRIGHT 2002 ELSEVIER SCI. B.V.DUPLICATE 1
ACCESSION NUMBER: 2000378102 EMBASE
TITLE: Thermus aquaticus DNA polymerase I mutants with altered fidelity. Interacting mutations in the
O-helix.
AUTHOR: Suzuki M.; Yoshida S.; Adman E.T.; Blank A.; Loeb L.A.; Gottstein J.
CORPORATE SOURCE: L.A. Loeb, J. Gottstein Mem. Cancer Res. Lab., Dept. of Pathology, University of Washington, Seattle, WA 98195-7705, United States. laloeb@u.washington.edu
SOURCE: Journal of Biological Chemistry, (20 Oct 2000) 275/42 (32728-32735).

Refs: 35
 ISN: 0021-9258 CODEN: JBCHA3
 COUNTRY: United States
 DOCUMENT TYPE: Journal; Article
 FILE SEGMENT: 029 Clinical Biochemistry
 LANGUAGE: English
 SUMMARY LANGUAGE: English
 AB **Phe667** in the conserved O-helix of *Thermus aquaticus* (Taq) DNA polymerase I (pol I) is known to be important for discrimination against dideoxy-NTPs. We show here that **Phe667** is also important for base selection fidelity. In a forward mutation assay at high polymerase concentration, wild type pol I catalyzed frequent A .fwdarw. T and G .fwdarw. T transversions and -1 frameshifts at nonreiterated sites involving loss of a purine immediately downstream of a pyrimidine. The mutants F667L and A661E,I665T, F667L exhibited large decreases in A .fwdarw. T and G .fwdarw. T transversions, and the triple mutant displayed reduction in the aforementioned -1 frameshifts as well. Kinetic analysis showed that the F667L and A661E,I665T, F667L polymerases discriminated against synthesis of A:A mispairs more effectively and catalyzed less extension of A:A mispairs than the wild type enzyme. These data indicate that **Phe667** functions in maintaining the error frequency and spectrum, and the catalytic efficiency, of wild type pol I. We also found that the strong general mutator activity conferred by the single A661E substitution was entirely suppressed in the A661E, I665T,F667L polymerase, exemplifying how interactions among O-helix residues can contribute to fidelity. We discuss the mutator and anti-mutator mutations in light of recently obtained three-dimensional structures of *T. aquaticus* pol I.

L3 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1999:571772 CAPLUS
 DOCUMENT NUMBER: 131:196102
 TITLE: Nucleotide compounds including a rigid linker
 INVENTOR(S): Khan, Shaheer H.; Rosenblum, Barnett B.; Zhen, Weiguo;
 Menchen, Steven M.
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S., 32 pp.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 5948648	A	19990907	US 1998-87250	19980529
US 6197555	B1	20010306	US 1999-461510	19991214
PRIORITY APPLN. INFO.:			US 1998-87250	A2 19980529
			US 1998-172789	A3 19981014

OTHER SOURCE(S): MARPAT 131:196102
 AB A nucleoside/tide compd. having a rigid linker attached to the 8-position of a purine, the 7-position of a 7-deazapurine and the 5-position of a pyrimidine is disclosed. Fluorescent dyes may be attached to this linker and the fluorescent nucleotide used in primer extension reactions. Thus, the fluorescein dye HEX-1 was attached to the 5-position of ddCTP via an acetylene-phenyl-oxyethyleneamino linkage. This nucleotide deriv. was used in DNA sequencing with Taq polymerase contg. an R660S mutation. The synthesis of a no. of nucleoside/nucleotide derivs. contg. various rigid linkers is described.

REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L3 ANSWER 4 OF 6 MEDLINE DUPLICATE 2
 ACCESSION NUMBER: 199907133 MEDLINE
 DOCUMENT NUMBER: 9907133 PubMed ID: 10101201
 TITLE: Dye structure affects Taq DNA polymerase terminator selectivity.
 AUTHOR: Brandis J W
 CORPORATE SOURCE: DNA Chemistry Group, Genetic Analysis Business Unit, PE Biosystems, 850 Lincoln Center Drive, Foster City, CA 94404, USA.. brandjw@perkin-elmer.com
 SOURCE: NUCLEIC ACIDS RESEARCH, (1999 Apr 15) 27 (8) 1912-8. Journal code: 08L; 0411011. ISSN: 0305-1048.
 PUB. COUNTRY: ENGLAND: United Kingdom
 LANGUAGE: English
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 199906
 ENTRY DATE: Entered STN: 19990628
 Last Updated on STN: 19990628
 Entered Medline: 19990614

AB All DNA sequencing methods have benefited from the development of new F667Y versions of Taq DNA polymerase. However, terminator chemistry methods show less uniform peak height patterns when compared to primer chemistry profiles suggesting that the dyes and/or their linker arms affect enzyme selectivity. We have measured elementary nucleotide rate and binding constants for representative rhodamine- and fluorescein-labeled terminators to determine how they interact with F667 versions of Taq Pol I. We have also developed a rapid gel-based selectivity assay that can be used to screen and to quantify dye-enzyme interactions with F667Y versions of the enzyme. Our results show that 6-TAMRA-ddTTP behaves like unlabeled ddTTP, while 6-FAM-ddTTP shows a 40-fold reduction in the rate constant for polymerization without affecting ground-state nucleotide binding. Detailed mechanism studies indicate that both isomers of different fluorescein dyes interfere with a conformational change step which the polymerase undergoes following nucleotide binding but only when these dyes are attached to pyrimidines. When these same dyes are attached to purines by the same propargylamino linker arm, they show no effect on enzyme selectivity. These studies suggest that it may be possible to develop fluorescein terminators for thermocycle DNA sequencing methods for polymerases that do not discriminate between deoxy- and dideoxynucleotides.

L3 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2002 ACS
 ACCESSION NUMBER: 1992:423036 CAPLUS
 DOCUMENT NUMBER: 117:23036
 TITLE: Regulation of tubercidin biosynthesis in Streptomyces tubercidicus by adenine and histidine
 AUTHOR(S): Yoo, Jin Cheol; Hah, Yung Chil
 CORPORATE SOURCE: Dep. Pharm., Chosun Univ., Kwangju, 501-759, S. Korea
 SOURCE: Misaengmul Hakhoechi (1991), 29(3), 160-6
 CODEN: MIHCAR; ISSN: 0440-2413
 DOCUMENT TYPE: Journal
 LANGUAGE: Korean
 AB The regulatory mechanism of tubercidin biosynthesis in S. tubercidicus was studied. In a wide-type strain, addn. of adenine and histidine into the medium decreased the tubercidin prodn. by 60-65% and 40%, resp. The effects of adenine and histidine were alleviated by the addn. of inosine monophosphate and 5-aminoimidazole-4-carboxamide ribotide. The prodn. of tubercidin in S. tubercidicus strain K115 (ade-) was nearly shut off by histidine. In contrast to strain K115, adenine inhibited tubercidin biosynthesis in S. tubercidicus strain K412 (his-). In S. tubercidicus F667 strain (ade-, his-), tubercidin prodn. was increased by adenine and histidine.

ACCESSION NUMBER: 1967:26509 CAPLUS

DOCUMENT NUMBER: 66:26509

TITLE: A mathematical method for the quantitative determination of proline and citrulline by automatic amino acid analysis

AUTHOR(S): Breuer, Josef; Ise, H.; Doellefeld, Erich; Breuer, Heinz

CORPORATE SOURCE: Abt. Klin. Chem. Biochem. Chirurgischen Univ. Poliklin., Bonn, Ger.

SOURCE: Z. Klin. Chem. (1966), 4(5), 267-8

CODEN: ZKLCAY

DOCUMENT TYPE: Journal

LANGUAGE: German

AB By using an automatic amino acid analyzer, it is possible to resolve mixts. of proline (I) and citrulline (II) by application of a math. method. The method is based on the differences in the absorption curves of the reaction products of ninhydrin and I and II, i.e., the I product has a broader absorption peak at 440 m.mu. than at 570 m.mu., while the reverse is true for II product. With norleucine as an internal standard, the following formulas were derived for the quant. detns.: I (mg.) = $(F440(KC570) - F570(KC440)) / (KP440(KC570) - KP570(KC440))$ and II (mg.) = $(F440(KP570) - F570(KP440)) / (KC440 - (KP570) - KC570(KP440))$, where F is the ratio of the area under the denoted wavelength absorption curve for test mixt. to the curve area for standard norleucine at 570 m.mu., and KC and KP are the curve areas for pure II

and

I, resp., at the denoted wavelengths.